

IFWP

RAW SEQUENCE LISTING DATE: 07/10/2006
PATENT APPLICATION: US/10/584,810 TIME: 10:24:35

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4 <110> APPLICANT: Merck & Co., Inc.
      6 <120> TITLE OF INVENTION: HCV RNA-DEPENDENT RNA POLYMERASE
     9 <130> FILE REFERENCE: 21471 PCT
C--> 11 <140> CURRENT APPLICATION NUMBER: US/10/584,810
C--> 11 <141> CURRENT FILING DATE: 2006-06-28
    11 <150> PRIOR APPLICATION NUMBER: 60/535,708
     12 <151> PRIOR FILING DATE: 2004-01-09
     14 <160> NUMBER OF SEQ ID NOS: 28
    16 <170> SOFTWARE: FastSEQ for Windows Version 4.0
    18 <210> SEQ ID NO: 1
     19 <211> LENGTH: 571
    20 <212> TYPE: PRT
     21 <213> ORGANISM: Artificial Sequence
    23 <220> FEATURE:
     24 <223> OTHER INFORMATION: modified HCV NS5B
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     29 Pro Glu Glu Lys Leu Pro Ile Asn Pro Leu Ser Asn Ser Leu Leu
                   20
                                        25
     31 Arg Tyr His Asn Lys Val Tyr Cys Thr Thr Thr Lys Ser Ala Ser Leu
                                   40
     33 Arg Ala Lys Lys Val Thr Phe Asp Arg Met Gln Val Leu Asp Ser Tyr
     35 Tyr Asp Ser Val Leu Lys Asp Ile Lys Leu Ala Ala Ser Lys Val Thr
                           70
     37 Ala Arg Leu Leu Thr Met Glu Glu Ala Cys Gln Leu Thr Pro Pro His
     39 Ser Ala Arg Ser Lys Tyr Gly Phe Gly Ala Lys Glu Val Arg Ser Leu
                                       105
     41 Ser Gly Arg Ala Val Asn His Ile Lys Ser Val Trp Lys Asp Leu Leu
                                   120
            115
    43 Glu Asp Ser Glu Thr Pro Ile Pro Thr Thr Ile Met Ala Lys Asn Glu
    44 130
                               135
     45 Val Phe Cys Val Asp Pro Thr Lys Gly Gly Lys Lys Ala Ala Arg Leu
                           150
                                                155
     47 Ile Val Tyr Pro Asp Leu Gly Val Arg Val Cys Glu Lys Met Ala Leu
                                           170
                       165
    49 Tyr Asp Ile Thr Gln Lys Leu Pro Gln Ala Val Met Gly Ala Ser Tyr
                   180
                                        185
    51 Gly Phe Gln Tyr Ser Pro Ala Gln Arg Val Glu Phe Leu Leu Lys Ala
                                   200
    53 Trp Ala Glu Lys Lys Asp Pro Met Gly Phe Ser Tyr Asp Thr Arg Cys
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54
      210
                          215
55 Phe Asp Ser Thr Val Thr Glu Arg Asp Ile Arg Thr Glu Glu Ser Ile
                      230
                                          235
57 Tyr Arg Ala Cys Ser Leu Pro Glu Glu Ala His Thr Ala Ile His Ser
                                      250
                  245
59 Leu Thr Glu Arg Leu Tyr Val Gly Pro Met Phe Asn Ser Lys Gly
                                  265
              260
61 Gln Thr Cys Gly Tyr Arg Arg Cys Arg Ala Ser Gly Val Leu Thr Thr
                              280
63 Ser Met Gly Asn Thr Ile Thr Cys Tyr Val Lys Ala Leu Ala Ala Cys
                          295
65 Lys Ala Ala Gly Ile Ile Ala Pro Thr Met Leu Val Cys Gly Asp Asp
                      310
                                          315
67 Leu Val Val Ile Ser Glu Ser Gln Gly Thr Glu Glu Asp Glu Arg Asn
                                      330
                  325
69 Leu Arg Ala Phe Thr Glu Ala Met Thr Arg Tyr Ser Ala Pro Pro Gly
              340
                                  345
71 Asp Pro Pro Arg Pro Glu Tyr Asp Leu Glu Leu Ile Thr Ser Cys Ser
                              360
73 Ser Asn Val Ser Val Ala Leu Gly Pro Gln Gly Arg Arg Tyr Tyr
                          375
75 Leu Thr Arg Asp Pro Thr Thr Pro Ile Ala Arg Ala Ala Trp Glu Thr
                      390
77 Val Arg His Ser Pro Val Asn Ser Trp Leu Gly Asn Ile Ile Gln Tyr
                                      410
                  405
79 Ala Pro Thr Ile Trp Ala Arg Met Val Leu Met Thr His Phe Phe Ser
                                  425
              420
81 Ile Leu Met Ala Gln Asp Thr Leu Asp Gln Asn Leu Asn Phe Glu Met
                              440
          435
83 Tyr Gly Ala Val Tyr Ser Val Ser Pro Leu Asp Leu Pro Ala Ile Ile
84 450
                          455
85 Glu Arg Leu His Gly Leu Asp Ala Phe Ser Leu His Thr Tyr Thr Pro
                      470
                                          475
87 His Glu Leu Thr Arq Val Ala Ser Ala Leu Arg Lys Leu Gly Ala Pro
                  485
                                      490
89 Pro Leu Arg Ala Trp Lys Ser Arg Ala Arg Ala Val Arg Ala Ser Leu
              500
                                  505
91 Ile Ser Arg Gly Gly Arg Ala Ala Val Cys Gly Arg Tyr Leu Phe Asn
          515
                              520
                                                   525
93 Trp Ala Val Lys Thr Lys Leu Lys Leu Thr Pro Leu Pro Glu Ala Arg
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                                              540
95 Leu Leu Asp Leu Ser Ser Trp Phe Thr Val Gly Ala Gly Gly Asp
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97 Ile Tyr His Ser Val Ser Arg Ala Arg Pro Arg
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101 <210> SEQ ID NO: 2
102 <211> LENGTH: 571
103 <212> TYPE: PRT
104 <213> ORGANISM: Artificial Sequence
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106 <220> FEATURE:																
107 <223> OTHER INFORMATION: modified HCV NS5B																
109 <400> SEQUENCE: 2 110 Met Ser Met Ser Tyr Thr Trp Thr Gly Ala Leu Ile Thr Pro Cys Gly																
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112	Pro	Glu	Glu	Glu	Lys	Leu	Pro	Ile	Asn	Pro	Leu	Ser	Asn	Ser	Leu	Met
113				20					25					30		
114	Arg	Phe	His	Asn	Lys	Val	Tyr	Ser	Thr	Thr	Ser	Arg	Ser	Ala	Ser	Leu
115			35					40					45		_	
116	Arg		Lys	Lys	Val	Thr		Asp	Arg	Val	Gln		Leu	Asp	Ala	His
117		50		_			55	_				60	_	_		_
	-	Asp	Ser	Val	Leu		Asp	Val	Lys	Arg		Ala	Ser	Lys	Val	
119		_	_	_		70	~-			_	75	_	1	_	.	80
	Ala	Arg	Leu	Leu		Val	GIu	GIu	Ala		Ala	Leu	Thr	Pro		HIS
121	_		_		85	_	~ 1	D 1	~1	90	T	~1	**- 7	7	95	T
	Ser	Ala	ьys		Arg	Tyr	GIY	Pne		Ата	гуѕ	GIU	vaı	Arg	ser	ьeu
123		•		100	*** 1	3	***	T 1 -	105	Q	T7_7	П	~1	110	T	T 0
	ser	Arg		Ala	vaı	ASI	HIS		Arg	ser	Val	пр	125	Asp	ьеи	Leu
125	a 1	7	115	TT	mb	Desc	т1.	120	mb ~	mb~	т1.	Mot		Trrc	7 cn	C111
		130	GIII	піѕ	TIIL	PLO	135	Asp	1111	1111	TIE	140	Ala	Lys	ASII	Gru
127			Cvc	Tla	λcn	Dro		Luc	G137	G1 ₃₇	Lvc		Dro	Ala	Δra	Len
	145	FIIE	Cys	116	vsh	150	1111	цуз	Gry	Сту	155	цуз	110	AIG	my	160
		Val	Тул	Pro	Δsn		Glv	Val	Δra	Val		Glu	Lvs	Met	Ala	
131	***	vai	- 7 -		165	шеш	011	val	3	170	O _I S	020	-1-		175	
	Tvr	Asp	Ile	Ala		Lvs	Leu	Pro	Lvs		Ile	Met	Glv	Pro		Tvr
133	-1-	P		180	V	-1-			185				1	190		4
	Glv	Phe	Gln		Ser	Pro	Ala	Glu		Val	Asp	Phe	Leu	Leu	Lys	Ala
135	•		195	-				200	_		_		205		_	
136	Trp	Gly	Ser	Lys	Lys	Asp	Pro	Met	Gly	Phe	Ser	Tyr	Asp	Thr	Arg	Cys
137	_	210					215					220				
138	Phe	Asp	Ser	Thr	Val	Thr	Glu	Arg	Asp	Ile	Arg	Thr	Glu	Glu	Ser	Ile
139	225					230					235					240
140	Tyr	Gln	Ala	Cys	Ser	Leu	Pro	Gln	Glu	Ala	Arg	Thr	Val	Ile	His	Ser
141					245					250					255	
142	Leu	Thr	Glu	Arg	Leu	Tyr	Val	Gly	Gly	Pro	Met	Thr	Asn	Ser	Lys	Gly
143				260					265			_		270	_	_
144	Gln	Ser		Gly	Tyr	Arg	Arg		Arg	Ala	Ser	Gly		Phe	Thr	Thr
145			275		_		_	280			_		285			_
	Ser		Gly	Asn	Thr	Met		Cys	Tyr	Ile	Lys		Leu	Ala	Ala	Cys
147	_	290				-	295	_			_	300	_	~3	_	_
	_	Ala	Ala	Gly	Ile		Asp	Pro	Val	Met		Val	Cys	Gly	Asp	
	305				_	310	_	~7	~7		315	~ 1	3	a 3	3	320
	Leu	Val	Val	lle		GIu	ser	Gin	GIY		GIU	GIU	Asp	Glu		ASI
151	.	7	7. T _	DI	325	a 7	70.7 -	Ma-	m\	330	M•	0	7.7 -	Dece	335	C1
	ьeu	arg	Ala		ınr	GIU	нта	мес		arg	ıyr	ser	AId	Pro 350	PLO	GIA
153	7	T	D	340	D====	~1	П	7	345	~1	T 0	т1 ^	Th∽		Cre	C.~
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Output Set: N:\CRF4\07102006\J584810.raw

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			Thr	Thr	Thr	Pro	Ile	Pro	Thr	Thr	Ile		Ala	Lys	Asn	Glu
210		130	C	** . 7	•	D	135	T	a 1	~ 1		140	D	77.	7	T
		Pne	Cys	vai	Asp	150	Ala	ьуѕ	GIY	GIY	_	ьys	Pro	Ата	Arg	160
	145	นาโ	Тъгъ	Dro	7 cn		Gly	v-1	λνα	17 a 1	155 Cvc	Glu	Luc	λνα	λ 1 =	
213	116	vai	ıyı	PIO	165	ьеи	GIY	vaı	Arg	170	Cys	Giu	цуз	Arg	175	Deu
	Tur	Δsn	Val	Tle		Lvs	Leu	Ser	Tle		Thr	Met	Glv	Ser		Tvr
216	- 7 -	1100	vai	180	01	 ,	DCu	501	185	014		1100		190		- 7 -
	Glv	Phe	Gln		Ser	Pro	Gln	Gln		Val	Glu	Ara	Leu		Lvs	Met
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220	-	210		-	-		215		_			220	_		_	_
221	Phe	Asp	Ser	Thr	Val	Thr	Glu	Gln	Asp	Ile	Arg	Val	Glu	Glu	Glu	Ile
222						230					235					240
223	Tyr	Gln	Cys	Cys	Asn	Leu	Glu	Pro	Glu	Ala	Arg	Lys	Val	Ile	Ser	Ser
224					245					250					255	
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226				260					265	_			_	270		_
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230	T	290	77.	a1	T	7	295	Dwa	7	Dha	T 011	300	C	~1	7 ~~	7 an
	-	Ата	Ата	GIY	ьeu	310	Asn	Pro	Asp	Pne		vai	Cys	GTÀ	Asp	320
	305	17 n 1	Wa l	17a l	λla		Ser	7 cn	Clar	17-1	315	Glu	λen	Δrα	Δla	
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236	LCu			340		014			345		-1-			350		1
	Asp	Ala	Pro		Pro	Thr	Tyr	Asp		Glu	Leu	Ile	Thr	Ser	Cys	Ser
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240		370					375					380				
241	Leu	Thr	Arg	Asp	Ala	Thr	Thr	Pro	Leu	Ala	Arg	Ala	Ala	${\tt Trp}$	Glu	Thr
242						390					395					400
243	Ala	Arg	His	Thr		Val	Asn	Ser	Trp		Gly	Asn	Ile	Ile		Tyr
244	_			_	405	_				410			•		415	_
	Ala	Pro	Thr		Trp	Val	Arg	Met		Met	Met	Thr	His		Phe	Ser
246		_	~7	420	~1	~7		_	425		D		.	430	a 1	3
	тте	Leu		Ser	Gin	GIU	Ile		Asp	Arg	Pro	ьeu		Pne	GIU	мет
248	(Tla ana	a 1	435	mb	m	C	77a 7	440	Desc	T 011	7 ~~	T 011	445	77.	т1.	T10
	Tyr		Ala	Int	Tyr	ser	Val	TIII	PIO	ьеи	Asp	460	PIO	Ala	iie	TTE
250	C1,,	450	T 011	Uic	C111	LOU	455 Ser	λla	Dho	Thr	Leu		Car	Тиг	Sar	Pro
251		AT 9	₽€U	1112	GIY	470	DET	та	FIIC	T 11T	475	1115	OCI	TYL	DCI	480
		Glu	Leu	Agn	Ara		Ala	Glv	Thr	Len		Lvs	Leu	Glv	Cvs	
254	V U.I.	Q1 u	Leu	21011	485	•41	111U	- 1		490	9	-,5		O-1	495	
	Pro	Leu	Ara	Ala		Ara	His	Ara	Ala		Ala	Val	Ara	Ala		Leu
256				500		د			505	J	- · · - - •			510	4	-
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VERIFICATION SUMMARY

NAME OF THE PARTY OF THE PARTY

DATE: 07/10/2006

PATENT APPLICATION: US/10/584,810

TIME: 10:24:36

Input Set: A:\21471 SEQ LSTG 6 16 06.TXT Output Set: N:\CRF4\07102006\J584810.raw

L:11 M:270 C: Current Application Number differs, Replaced Current Application No

L:11 M:271 C: Current Filing Date differs, Replaced Current Filing Date